

## TENT COOPERATION TREA.

From the INTERNATIONAL BUREAU

PCT

## NOTIFICATION OF ELECTION

(PCT Rule 61.2)

Date of mailing (day/month/year) 03 July 2001 (03.07.01)
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To:  Commissioner US Department of Commerce United States Patent and Trademark Office, PCT 2011 South Clark Place Room CP2/5C24 Arlington, VA 22202 ETATS-UNIS D'AMERIQUE  in its capacity as elected Office
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International application No. PCT/EP00/08487
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Applicant's or agent's file reference PU3517
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International filing date (day/month/year) 31 August 2000 (31.08.00)
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Priority date (day/month/year) 04 September 1999 (04.09.99)
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Applicant
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PIANETTI, Pascal, Maurice, Charles et al
--

1. The designated Office is hereby notified of its election made:

in the demand filed with the International Preliminary Examining Authority on:

23 March 2001 (23.03.01)

in a notice effecting later election filed with the International Bureau on:

2. The election  was

was not

made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b).

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland
---

Authorized officer
--------------------

Zakaria EL KHODARY
--------------------

Facsimile No.: (41-22) 740.14.35
----------------------------------

Telephone No.: (41-22) 338.83.38
----------------------------------

## PATENT COOPERATION TREATY

PCT

NOTIFICATION OF THE RECORDING  
OF A CHANGE(PCT Rule 92bis.1 and  
Administrative Instructions, Section 422)Date of mailing (day/month/year)  
13 juillet 2001 (13.07.01)

From the INTERNATIONAL BUREAU

To:

CRAWLEY, Karen  
GlaxoSmithKline  
Corporate Intellectual Property  
Two New Horizons Court  
Brentford  
Middlesex TW8 9EP  
ROYAUME-UNIApplicant's or agent's file reference  
PU3517

## IMPORTANT NOTIFICATION

International application No.  
PCT/EP00/08487International filing date (day/month/year)  
31 août 2000 (31.08.00)

1. The following indications appeared on record concerning:

 the applicant  the inventor  the agent  the common representative

## Name and Address

CRAWLEY, Karen  
Glaxo Wellcome plc  
Berkeley Avenue  
Greenford, Middlesex UB6 0NN  
United Kingdom

State of Nationality

State of Residence

Telephone No.

020 8966 8000

Facsimile No.

020 8966 8838

Teleprinter No.

2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

 the person  the name  the address  the nationality  the residence

## Name and Address

CRAWLEY, Karen  
GlaxoSmithKline  
Corporate Intellectual Property  
Two New Horizons Court  
Brentford  
Middlesex TW8 9EP  
United Kingdom

State of Nationality

State of Residence

Telephone No.

020 8966 8412

Facsimile No.

020 8966 8838

Teleprinter No.

3. Further observations, if necessary:

4. A copy of this notification has been sent to:

 the receiving Office the designated Offices concerned the International Searching Authority the elected Offices concerned the International Preliminary Examining Authority other:The International Bureau of WIPO  
34, chemin des Colombettes  
1211 Geneva 20, Switzerland

Authorized officer

Christine Carrié

Facsimile No.: (41-22) 740.14.35

Telephone No.: (41-22) 338.83.38

## INTERNATIONAL COOPERATION TREATY

PCT

NOTIFICATION OF THE RECORDING  
OF A CHANGE(PCT Rule 92bis.1 and  
Administrative Instructions, Section 422)

Date of mailing (day/month/year) 14 February 2002 (14.02.02)		From the INTERNATIONAL BUREAU																		
Applicant's or agent's file reference PU3517		To: CRAWLEY, Karen GlaxoSmithKline Corporate Intellectual Property (CN9.25.1) 980 Great West Road Brentford Middlesex TW8 9GS ROYAUME-UNI																		
International application No. PCT/EP00/08487		IMPORTANT NOTIFICATION																		
		International filing date (day/month/year) 31 August 2000 (31.08.00)																		
<p>1. The following indications appeared on record concerning:</p> <p><input type="checkbox"/> the applicant    <input type="checkbox"/> the inventor    <input checked="" type="checkbox"/> the agent    <input type="checkbox"/> the common representative</p> <table border="1"> <tr> <td rowspan="4">Name and Address  CRAWLEY, Karen GlaxoSmithKline Corporate Intellectual Property Two New Horizons Court Brentford Middlesex TW8 9EP United Kingdom</td> <td colspan="2">State of Nationality</td> <td colspan="2">State of Residence</td> </tr> <tr> <td colspan="2">Telephone No. 020 8966 8412</td> <td colspan="2"></td> </tr> <tr> <td colspan="2">Facsimile No. 020 8966 8838</td> <td colspan="2"></td> </tr> <tr> <td colspan="2">Teleprinter No.</td> <td colspan="2"></td> </tr> </table>				Name and Address  CRAWLEY, Karen GlaxoSmithKline Corporate Intellectual Property Two New Horizons Court Brentford Middlesex TW8 9EP United Kingdom	State of Nationality		State of Residence		Telephone No. 020 8966 8412				Facsimile No. 020 8966 8838				Teleprinter No.			
Name and Address  CRAWLEY, Karen GlaxoSmithKline Corporate Intellectual Property Two New Horizons Court Brentford Middlesex TW8 9EP United Kingdom	State of Nationality		State of Residence																	
	Telephone No. 020 8966 8412																			
	Facsimile No. 020 8966 8838																			
	Teleprinter No.																			
<p>2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:</p> <p><input type="checkbox"/> the person    <input type="checkbox"/> the name    <input checked="" type="checkbox"/> the address    <input type="checkbox"/> the nationality    <input type="checkbox"/> the residence</p> <table border="1"> <tr> <td rowspan="4">Name and Address  CRAWLEY, Karen GlaxoSmithKline Corporate Intellectual Property (CN9.25.1) 980 Great West Road Brentford Middlesex TW8 9GS United Kingdom</td> <td colspan="2">State of Nationality</td> <td colspan="2">State of Residence</td> </tr> <tr> <td colspan="2">Telephone No. 44 20 8047 5000</td> <td colspan="2"></td> </tr> <tr> <td colspan="2">Facsimile No. 44 20 8047 6894</td> <td colspan="2"></td> </tr> <tr> <td colspan="2">Teleprinter No.</td> <td colspan="2"></td> </tr> </table>				Name and Address  CRAWLEY, Karen GlaxoSmithKline Corporate Intellectual Property (CN9.25.1) 980 Great West Road Brentford Middlesex TW8 9GS United Kingdom	State of Nationality		State of Residence		Telephone No. 44 20 8047 5000				Facsimile No. 44 20 8047 6894				Teleprinter No.			
Name and Address  CRAWLEY, Karen GlaxoSmithKline Corporate Intellectual Property (CN9.25.1) 980 Great West Road Brentford Middlesex TW8 9GS United Kingdom	State of Nationality		State of Residence																	
	Telephone No. 44 20 8047 5000																			
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	Teleprinter No.																			
<p>3. Further observations, if necessary:</p>																				
<p>4. A copy of this notification has been sent to:</p> <table> <tr> <td><input checked="" type="checkbox"/> the receiving Office</td> <td><input type="checkbox"/> the designated Offices concerned</td> </tr> <tr> <td><input type="checkbox"/> the International Searching Authority</td> <td><input checked="" type="checkbox"/> the elected Offices concerned</td> </tr> <tr> <td><input type="checkbox"/> the International Preliminary Examining Authority</td> <td><input type="checkbox"/> other:</td> </tr> </table>				<input checked="" type="checkbox"/> the receiving Office	<input type="checkbox"/> the designated Offices concerned	<input type="checkbox"/> the International Searching Authority	<input checked="" type="checkbox"/> the elected Offices concerned	<input type="checkbox"/> the International Preliminary Examining Authority	<input type="checkbox"/> other:											
<input checked="" type="checkbox"/> the receiving Office	<input type="checkbox"/> the designated Offices concerned																			
<input type="checkbox"/> the International Searching Authority	<input checked="" type="checkbox"/> the elected Offices concerned																			
<input type="checkbox"/> the International Preliminary Examining Authority	<input type="checkbox"/> other:																			
The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland		Authorized officer Sean Taylor																		
Facsimile No.: (41-22) 740.14.35		Telephone No.: (41-22) 338.83.38																		

## PATENT COOPERATION TREATY

10/07/084

PCT

NOTIFICATION OF THE RECORDING  
OF A CHANGE(PCT Rule 92bis.1 and  
Administrative Instructions, Section 422)

Date of mailing (day/month/year) 10 décembre 2002 (10.12.02)	From the INTERNATIONAL BUREAU To:  CRAWLEY, Karen GlaxoSmithKline Corporate Intellectual Property (CN9.25.1) 980 Great West Road Brentford Middlesex TW8 9GS United Kingdom	
Applicant's or agent's file reference PU3517	IMPORTANT NOTIFICATION	
International application No. PCT/EP00/08487	International filing date (day/month/year) 31 août 2000 (31.08.00)	

## 1. The following indications appeared on record concerning:

the applicant  the inventor  the agent  the common representative

Name and Address PIANETTI, Pascal, Maurice, Charles	State of Nationality FR	State of Residence FR
	Telephone No.	
	Facsimile No.	
	Teleprinter No.	

## 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

the person  the name  the address  the nationality  the residence

Name and Address PIANETTI, Pascal, Maurice, Charles Laboratoire GlaxoSmithKline Centre de Recherches Z.A. de Courtabœuf 25, Avenue de Québec F-91940 Les Ulis France	State of Nationality FR	State of Residence FR
	Telephone No.	
	Facsimile No.	
	Teleprinter No.	

## 3. Further observations, if necessary:

Corrected version of IB/306 mailed on 26 March 2002. The person in box No. 2 will be the applicant/inventor for US only.

## 4. A copy of this notification has been sent to:

<input checked="" type="checkbox"/> the receiving Office	<input type="checkbox"/> the designated Offices concerned
<input type="checkbox"/> the International Searching Authority	<input checked="" type="checkbox"/> the elected Offices concerned
<input checked="" type="checkbox"/> the International Preliminary Examining Authority	<input type="checkbox"/> other:

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland  Facsimile No.: (41-22) 740.14.35	Authorized officer Ki-Nam HA  Telephone No.: (41-22) 338.83.38
---	---

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TECH CENTER 1600/2900  
FEB 12 2003

## PATENT COOPERATION TREATY

PCT

NOTIFICATION OF THE RECORDING  
OF A CHANGE(PCT Rule 92bis.1 and  
Administrative Instructions, Section 422)Date of mailing (day/month/year)  
26 March 2002 (26.03.02)

## From the INTERNATIONAL BUREAU

To:

CRAWLEY, Karen  
GlaxoSmithKline  
Corporate Intellectual Property  
(CN9.25.1)  
980 Great West Road  
Brentford  
Middlesex TW8 9GS  
ROYAUME-UNIApplicant's or agent's file reference  
PU3517

## IMPORTANT NOTIFICATION

International application No.  
PCT/EP00/08487International filing date (day/month/year)  
31 August 2000 (31.08.00)

## 1. The following indications appeared on record concerning:

 the applicant     the inventor     the agent     the common representative
Name and Address  
PIANETTI, Pascal, Maurice, Charles  
Laboratoire Glaxo Wellcome  
Centre de Recherches  
25, avenue de Québec  
F-91951 Les Ulis  
France

State of Nationality FR	State of Residence FR
Telephone No.	
Facsimile No.	
Teleprinter No.	

## 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

 the person     the name     the address     the nationality     the residence
Name and Address  
PIANETTI, Pascal, Maurice, Charles  
Laboratoire Glaxo Wellcome  
Centre de Recherches  
25, avenue de Québec  
F-91951 Les Ulis  
France

State of Nationality FR	State of Residence FR
Telephone No.	
Facsimile No.	
Teleprinter No.	

## 3. Further observations, if necessary:

The person in box No. 2 will be the applicant for all designated States except US.

## 4. A copy of this notification has been sent to:

<input checked="" type="checkbox"/> the receiving Office	<input type="checkbox"/> the designated Offices concerned
<input type="checkbox"/> the International Searching Authority	<input checked="" type="checkbox"/> the elected Offices concerned
<input type="checkbox"/> the International Preliminary Examining Authority	<input type="checkbox"/> other:

The International Bureau of WIPO 34, chemin des Colombelettes 1211 Geneva 20, Switzerland Facsimile No.: (41-22) 740.14.35	Authorized officer Ki-Nam HA Telephone No.: (41-22) 338.83.38
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## PENT COOPERATION TREA

PCT

NOTIFICATION OF THE RECORDING  
OF A CHANGE(PCT Rule 92bis.1 and  
Administrative Instructions, Section 422)

Date of mailing (day/month/year) 13 March 2001 (13.03.01)	
Applicant's or agent's file reference PU3517	IMPORTANT NOTIFICATION
International application No. PCT/EP00/08487	International filing date (day/month/year) 31 August 2000 (31.08.00)

## 1. The following indications appeared on record concerning:

the applicant  the inventor  the agent  the common representative

Name and Address PIANELLI, Pascal, Maurice, Charles Laboratoire Glaxo Wellcome Centre de Recherches Z.A. de Courabœuf 25, avenue de Québec F-91940 Les Ulis France	State of Nationality FR	State of Residence FR
	Telephone No.	
	Facsimile No.	
	Teleprinter No.	

## 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

the person  the name  the address  the nationality  the residence

Name and Address PIANELLI, Pascal, Maurice, Charles Laboratoire Glaxo Wellcome Centre de Recherches 25, avenue de Québec F-91951 Les Ulis France	State of Nationality FR	State of Residence FR
	Telephone No.	
	Facsimile No.	
	Teleprinter No.	

## 3. Further observations, if necessary:

**The Applicant's status has been changed into applicant/inventor for the US only.**

## 4. A copy of this notification has been sent to:

<input checked="" type="checkbox"/> the receiving Office	<input checked="" type="checkbox"/> the designated Offices concerned
<input type="checkbox"/> the International Searching Authority	<input type="checkbox"/> the elected Offices concerned
<input type="checkbox"/> the International Preliminary Examining Authority	<input type="checkbox"/> other:

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Facsimile No.: (41-22) 740.14.35	Authorized officer Eugénia Santos Telephone No.: (41-22) 338.83.38
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## PATENT COOPERATION TREATY

PCT

NOTIFICATION OF THE RECORDING  
OF A CHANGE(PCT Rule 92bis.1 and  
Administrative Instructions, Section 422)

Date of mailing (day/month/year) 19 April 2002 (19.04.02)
Applicant's or agent's file reference PU3517
International application No. PCT/EP00/08487

## From the INTERNATIONAL BUREAU

To:

CRAWLEY, Karen  
GlaxoSmithKline  
Corporate Intellectual Property  
(CN9.25.1)  
980 Great West Road  
Brentford  
Middlesex TW8 9GS  
ROYAUME-UNI

## IMPORTANT NOTIFICATION

International filing date (day/month/year) 31 August 2000 (31.08.00)
---

## 1. The following indications appeared on record concerning:

the applicant  the inventor  the agent  the common representative

## Name and Address

PIANETTI, Pascal, Maurice, Charles  
Laboratoire Glaxo Wellcome  
Centre de Recherches  
25, avenue de Québec  
F-91951 Les Ulis  
France

## State of Nationality

FR

## State of Residence

FR

Telephone No.

Facsimile No.

Teleprinter No.

## 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

the person  the name  the address  the nationality  the residence

## Name and Address

PIANETTI, Pascal, Maurice, Charles  
Laboratoire GlaxoSmithKline  
Centre de Recherches  
Z.A. de Courabœuf  
25, Avenue de Québec  
F-91940 Les Ulis  
France

## State of Nationality

FR

## State of Residence

FR

Telephone No.

Facsimile No.

Teleprinter No.

## 3. Further observations, if necessary:

## 4. A copy of this notification has been sent to:

the receiving Office

the designated Offices concerned

the International Searching Authority

the elected Offices concerned

the International Preliminary Examining Authority

other:

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland  Facsimile No.: (41-22) 740.14.35	Authorized officer  Kiwa MPAY  Telephone No.: (41-22) 338.83.38
---	---

# INTERNATIONAL SEARCH REPORT

Inten [REDACTED] Application No  
PCT/EP/00/08487

A. CLASSIFICATION OF SUBJECT MATTER				
IPC 7	C07D277/24	A61K31/16	A61K31/33	A61P31/00
	C07D307/80	C07D333/56	C07D207/32	C07D409/04
	C07C311/46	C07D233/22	C07D295/08	C07D277/62

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 C07D C07C A61K A61P

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, CHEM ABS Data, BEILSTEIN Data

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	P G WYATT ET AL: "JOURNAL OF MEDICINAL CHEMISTRY, US, AMERICAN CHEMICAL SOCIETY. WASHINGTON" JOURNAL OF MEDICINAL CHEMISTRY, US, AMERICAN CHEMICAL SOCIETY. WASHINGTON, vol. 38, no. 10, 1995, pages 1657-1665, XP002084134 ISSN: 0022-2623 page 1658; table 1 page 1659; table 2 page 1660; table 3	1-37
P, X	WO 99 55682 A (UNIV GEORGETOWN) 4 November 1999 (1999-11-04) figure 10	1

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

### \* Special categories of cited documents :

- \*A\* document defining the general state of the art which is not considered to be of particular relevance
- \*E\* earlier document but published on or after the international filing date
- \*L\* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- \*O\* document referring to an oral disclosure, use, exhibition or other means
- \*P\* document published prior to the international filing date but later than the priority date claimed

- \*T\* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- \*X\* document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- \*Y\* document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- \*8\* document member of the same patent family

Date of the actual completion of the international search

6 February 2001

Date of mailing of the international search report

21/02/2001

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2  
NL - 2280 HV Rijswijk  
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl.  
Fax: (+31-70) 340-3016

Authorized officer

Pauwels, G

# INTERNATIONAL SEARCH REPORT

Int. Application No.  
PCT/EP 00/08487

## A. CLASSIFICATION OF SUBJECT MATTER

IPC 7	C07D277/64	C07D275/04	C07D233/84	C07D277/36	C07D277/28
	C07D263/32	C07C317/40	C07D521/00	C07D285/06	C07D231/56
	C07D409/12	C07D333/22	C07D241/44	C07D213/50	C07D235/28

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	FR 1 552 793 A (ETS. CLYN-BILA) 10 January 1969 (1969-01-10) page 6; example 56	1
X	US 4 883 815 A (ASCHWENDEN WERNER ET AL) 28 November 1989 (1989-11-28) example 27	1
X	US 4 207 234 A (HASHIMOTO MASASHI ET AL) 10 June 1980 (1980-06-10) example 190	1
X	SUGASAWA, TSUTOMU ET AL: "1-Azacycloalkyl-1,4-benzodiazepin-2-ones with antianxiety- antidepressant actions" J. MED. CHEM. (1985), 28(6), 699-707 XP002159556 page 700; figure 15	1
		-/-

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

### \* Special categories of cited documents :

- \*A\* document defining the general state of the art which is not considered to be of particular relevance
- \*E\* earlier document but published on or after the international filing date
- \*L\* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- \*O\* document referring to an oral disclosure, use, exhibition or other means
- \*P\* document published prior to the international filing date but later than the priority date claimed

- \*T\* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- \*X\* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- \*Y\* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- \*&\* document member of the same patent family

Date of the actual completion of the international search

6 February 2001

Date of mailing of the international search report

Name and mailing address of the ISA  
European Patent Office, P.B. 5818 Patentlaan 2  
NL - 2280 HV Rijswijk  
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  
Fax: (+31-70) 340-3016

Authorized officer

Pauwels, G

# INTERNATIONAL SEARCH REPORT

Intern. Application No.  
PCT/EP 00/08487

## A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 C07D235/06 C07D235/26 C07D209/08 C07C259/06 C07C235/24  
 C07D295/12 C07D249/18 C07D233/54 C07D333/38 C07D213/61  
 C07D401/12 C07C323/22 C07C317/24 C07C311/24 C07D213/64

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	CAPUANO, LILLY ET AL: "Darstellung von alpha-'2- alkoxy carbonyl-anilino!- und alpha-'2-Acylanilino!-carbonsäureamiden" JUSTUS LIEBIGS ANN. CHEM. (1968), 712, 73-8 XP002159557 page 73; figure 4J page 78, paragraph 1	1
X	BRANDL, E.: "Penicillin reactions. 1. Behavior towards penicillinase" SCI. PHARM. (1971), 39(4), 267-92 XP000979965 page 279; example 95	1

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

### \* Special categories of cited documents :

- \*A\* document defining the general state of the art which is not considered to be of particular relevance
- \*E\* earlier document but published on or after the international filing date
- \*L\* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other, special reason (as specified)
- \*O\* document referring to an oral disclosure, use, exhibition or other means
- \*P\* document published prior to the international filing date but later than the priority date claimed

- \*T\* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- \*X\* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- \*Y\* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- \*&\* document member of the same patent family

Date of the actual completion of the international search

6 February 2001

Date of mailing of the international search report

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2  
 NL - 2280 HV Rijswijk  
 Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  
 Fax: (+31-70) 340-3016

Authorized officer

Pauwels, G

# INTERNATIONAL SEARCH REPORT

Interr. Application No  
PCT/EP 00/08487

## A. CLASSIFICATION OF SUBJECT MATTER

IPC 7	C07D333/72	C07D215/08	C07D217/06	C07D213/75	C07D261/16
	C07C311/08	C07D211/44	C07D407/12	C07D307/46	C07D333/54
	C07D213/42	C07D333/20	C07D207/09	C07D307/14	C07D207/20

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

### \* Special categories of cited documents :

- \*A\* document defining the general state of the art which is not considered to be of particular relevance
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- \*L\* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- \*O\* document referring to an oral disclosure, use, exhibition or other means
- \*P\* document published prior to the international filing date but later than the priority date claimed

- \*T\* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- \*X\* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- \*Y\* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- \*Z\* document member of the same patent family

Date of the actual completion of the international search

6 February 2001

Date of mailing of the international search report

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2  
NL - 2280 HV Rijswijk  
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl  
Fax (+31-70) 340-3016

Authorized officer

Pauwels, G

# INTERNATIONAL SEARCH REPORT

Intell. Application No  
PCT/EP 00/08487

**A. CLASSIFICATION OF SUBJECT MATTER**  
 IPC 7 C07D213/85 C07D213/84 C07D333/32 C07C255/56 C07C275/10  
 C07D295/22 C07C255/44 C07D295/14 C07C237/42

According to International Patent Classification (IPC) or to both national classification and IPC

**B. FIELDS SEARCHED**

Minimum documentation searched (classification system followed by classification symbols)

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

**C. DOCUMENTS CONSIDERED TO BE RELEVANT**

Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

\* Special categories of cited documents :

- \*A\* document defining the general state of the art which is not considered to be of particular relevance
- \*E\* earlier document but published on or after the international filing date
- \*L\* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- \*O\* document referring to an oral disclosure, use, exhibition or other means
- \*P\* document published prior to the international filing date but later than the priority date claimed

- \*T\* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- \*X\* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- \*Y\* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- \*&\* document member of the same patent family

Date of the actual completion of the international search

Date of mailing of the international search report

6 February 2001

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patenttaan 2  
NL - 2280 HV Rijswijk  
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl.  
Fax: (+31-70) 340-3016

Authorized officer

Pauwels, G

**INTERNATIONAL SEARCH REPORT**

Information on patent family members

Interr.	Application No
PCT/EP 00/08487	

Patent document cited in search report		Publication date		Patent family member(s)		Publication date
WO 9955682	A	04-11-1999	AU	3669199 A		16-11-1999
FR 1552793	A	10-01-1969	DE	1643317 A		16-06-1971
			GB	1197597 A		08-07-1970
			US	3551478 A		29-12-1970
US 4883815	A	28-11-1989	AU	603470 B		15-11-1990
			AU	1917688 A		27-01-1989
			CN	1036952 A, B		08-11-1989
			DD	277071 A		21-03-1990
			DK	405888 A		22-01-1989
			EP	0300371 A		25-01-1989
			FI	883441 A		22-01-1989
			GB	2207135 A, B		25-01-1989
			HU	47077 A, B		30-01-1989
			IN	167395 A		20-10-1990
			JP	1047748 A		22-02-1989
			MC	1960 A		30-06-1989
			MT	1023 A		18-07-1989
			MW	2688 A		12-04-1989
			NO	883233 A		23-01-1989
			NZ	225442 A		28-05-1991
			OA	8746 A		31-03-1989
			PT	88041 A		30-06-1989
			US	5017608 A		21-05-1991
			YU	39890 A		31-10-1991
			YU	141488 A		30-06-1990
			ZA	8805102 A		29-03-1989
			ZW	9288 A		01-02-1989
US 4207234	A	10-06-1980	US	4472300 A		18-09-1984

## PATENT COOPERATION TREATY

23 MAR 2001	PCT
NOTIFICATION OF THE RECORDING OF A CHANGE	

(PCT Rule 92bis.1 and  
Administrative Instructions, Section 422)

Date of mailing (day/month/year) 13 March 2001 (13.03.01)
Applicant's or agent's file reference PU3517
International application No. PCT/EP00/08487

From the INTERNATIONAL BUREAU

To:

CRAWLEY, Karen  
Glaxo Wellcome plc  
Berkeley Avenue  
Greenford, Middlesex UB6 0NN  
ROYAUME-UNI

## IMPORTANT NOTIFICATION

International filing date (day/month/year)  
31 August 2000 (31.08.00)

## 1. The following indications appeared on record concerning:

the applicant  the inventor  the agent

the common representative

Name and Address  
PIANETTI, Pascal, Maurice, Charles  
Laboratoire Glaxo Wellcome  
Centre de Recherches  
Z.A. de Courabœuf  
25, avenue de Québec  
F-91940 Les Ulis  
France

State of Nationality  
FR

State of Residence  
FR

Telephone No.

Facsimile No.

Teleprinter No.

## 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

the person  the name  the address  the nationality  the residence

Name and Address  
PIANETTI, Pascal, Maurice, Charles  
Laboratoire Glaxo Wellcome  
Centre de Recherches  
25, avenue de Québec  
F-91951 Les Ulis  
France

State of Nationality  
FR

State of Residence  
FR

Telephone No.

Facsimile No.

Teleprinter No.

## 3. Further observations, if necessary:

The Applicant's status has been changed into applicant/inventor for the US only.

## 4. A copy of this notification has been sent to:

the receiving Office  
 the International Searching Authority  
 the International Preliminary Examining Authority

the designated Offices concerned  
 the elected Offices concerned  
 other:

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Facsimile No.: (41-22) 740.14.35	Authorized officer Eugénia Santos Telephone No.: (41-22) 338.83.38
---	--

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## PATENT COOPERATION TREATY

RECEIVED  
23 JUL 2001  
NEW HORIZONS  
PCT

NOTIFICATION OF THE RECORDING  
OF A CHANGE

(PCT Rule 92bis.1 and  
Administrative Instructions, Section 422)

Date of mailing (day/month/year)  
13 July 2001 (13.07.01)

Applicant's or agent's file reference  
PU3517

International application No.  
PCT/EP00/08487

From the INTERNATIONAL BUREAU

To:

CRAWLEY, Karen  
GlaxoSmithKline  
Corporate Intellectual Property  
Two New Horizons Court  
Brentford  
Middlesex TW8 9EP  
ROYAUME-UNI

## IMPORTANT NOTIFICATION

International filing date (day/month/year)  
31 August 2000 (31.08.00)

## 1. The following indications appeared on record concerning:

the applicant  the inventor  the agent  the common representative

## Name and Address

CRAWLEY, Karen  
Glaxo Wellcome plc  
Berkeley Avenue  
Greenford, Middlesex  
United Kingdom

Corporate Intellectual Property	
RECEIVED	UNIT
B6 0NN	
24 JUL 2001	
OK	
Action Date	ATTN
	FILE

## State of Nationality State of Residence

Telephone No.  
020 8966 8000  
Facsimile No.  
020 8966 8838  
Teleprinter No.

## 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

the person  the name  the address  the nationality  the residence

## Name and Address

CRAWLEY, Karen  
GlaxoSmithKline  
Corporate Intellectual Property  
Two New Horizons Court  
Brentford  
Middlesex TW8 9EP  
United Kingdom

State of Nationality State of Residence

Telephone No.  
020 8966 8412  
Facsimile No.  
020 8966 8838  
Teleprinter No.

## 3. Further observations, if necessary:

## 4. A copy of this notification has been sent to:

the receiving Office  the designated Offices concerned  
 the International Searching Authority  the elected Offices concerned  
 the International Preliminary Examining Authority  other:

The International Bureau of WIPO  
34, chemin des Colombettes  
1211 Geneva 20, Switzerland  
Facsimile No.: (41-22) 740.14.35

Authorized officer

*e. carrie*  
Christine Carrie

Telephone No.: (41-22) 338.83.38

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PCT REQUEST

Original (for SUBMISSION) - printed on 15.08.2000 04:08:03 PM

PU3517

0-1	For receiving Office use only International Application No.	<b>PCT/EP 00/08487</b>	
0-2	International Filing Date	31 AUG 2000 - (31 08 2000)	
0-3	Name of receiving Office and "PCT International Application"	EUROPEAN PATENT OFFICE PCT INTERNATIONAL APPLICATION	
0-4 0-4-1	Form - PCT/RO/101 PCT Request Prepared using	PCT-EASY Version 2.91 (updated 01.07.2000)	
0-5	Petition The undersigned requests that the present international application be processed according to the Patent Cooperation Treaty		
0-6	Receiving Office (specified by the applicant)	European Patent Office (EPO) (RO/EP)	
0-7	Applicant's or agent's file reference	PU3517	
I	Title of Invention	BENZOPHENONES AS INHIBITORS OF REVERSE TRANSCRIPTASE	
II	Applicant This person is:	applicant only	
II-1		all designated States, except US	
II-2	Applicant for	GLAXO GROUP LIMITED	
II-4	Name	Glaxo Wellcome House	
II-5	Address:	Berkeley Avenue	
II-6	State of nationality	Greenford, Middlesex UB6 0NN	
II-7	State of residence	United Kingdom	
II-8	Telephone No.	GB	
II-9	Fax/Email No.	020 8966 8000 020 8966 8838	
III-1 III-1-1	Applicant and/or inventor This person is:	applicant and inventor	
III-1-2	Applicant for	US only	
III-1-4	Name (LAST, First)	ANDREWS, Clarence, Webster	
III-1-5	Address:	Glaxo Wellcome Inc. Five Moore Drive	
III-1-6	State of nationality	Research Triangle Park, NC NC 27709	
III-1-7	State of residence	United States of America US US	

## PCT REQUEST

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PU3517

III-2	Applicant and/or inventor	
III-2-1	This person is:	applicant and inventor
III-2-2	Applicant for	US only
III-2-4	Name (LAST, First)	CHAN, Joseph, Howing
III-2-5	Address:	Glaxo Wellcome Inc Five Moore Drive Research Triangle Park, NC NC 27709
III-2-6	State of nationality	United States of America
III-2-7	State of residence	US
III-3	Applicant and/or inventor	
III-3-1	This person is:	applicant and inventor
III-3-2	Applicant for	US only
III-3-4	Name (LAST, First)	FREEMAN, George, Andrew
III-3-5	Address:	Glaxo Wellcome Inc. Five Moore Drive Research Triangle Park, NC NC 27709
III-3-6	State of nationality	United States of America
III-3-7	State of residence	US
III-4	Applicant and/or inventor	
III-4-1	This person is:	applicant only
III-4-2	Applicant for	all designated States except US
III-4-4	Name (LAST, First)	PIANETTI, Pascal, Maurice, Charles
III-4-5	Address:	Laboratoire Glaxo Wellcome Centre de Recherches Z.A. de Courabœuf F-91940 Les Ulis 25 avenue de Quebec
III-4-6	State of nationality	France
III-4-7	State of residence	FR
III-5	Applicant and/or inventor	
III-5-1	This person is:	applicant and inventor
III-5-2	Applicant for	US only
III-5-4	Name (LAST, First)	ROMINES, Karen, Rene
III-5-5	Address:	Glaxo Wellcome Inc Five Moore Drive Research Triangle Park, NC NC 27709
III-5-6	State of nationality	United States of America
III-5-7	State of residence	US

## PCT REQUEST

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PU3517

III-6	Applicant and/or inventor	
III-6-1	This person is:	
III-6-2	Applicant for	
III-6-4	Name (LAST, First)	
III-6-5	Address:	
III-6-6	State of nationality	
III-6-7	State of residence	
III-7	Applicant and/or Inventor	
III-7-1	This person is:	
III-7-2	Applicant for	
III-7-4	Name (LAST, First)	
III-7-5	Address:	
III-7-6	State of nationality	
III-7-7	State of residence	
IV-1	Agent or common representative; or address for correspondence	
The person identified below is hereby/has been appointed to act on behalf of the applicant(s) before the competent International Authorities as:		
IV-1-1	Name (LAST, First)	
IV-1-2	Address:	
IV-1-3	Telephone No.	
IV-1-4	Facsimile No.	

## PCT REQUEST

4/5

Original (for SUBMISSION) - printed on 15.08.2000 04:08:03 PM

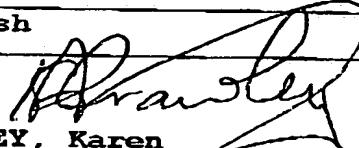
PU3517

V	Designation of States	
V-1	Regional Patent (other kinds of protection or treatment, if any, are specified between parentheses after the designation(s) concerned)	<p>AP: GH GM KE LS MW MZ SD SL SZ TZ UG ZW and any other State which is a Contracting State of the Harare Protocol and of the PCT</p> <p>EA: AM AZ BY KG KZ MD RU TJ TM and any other State which is a Contracting State of the Eurasian Patent Convention and of the PCT</p> <p>EP: AT BE CH&amp;LI CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE and any other State which is a Contracting State of the European Patent Convention and of the PCT</p> <p>OA: BF BJ CF CG CI CM GA GN GW ML MR NE SN TD TG and any other State which is a member State of OAPI and a Contracting State of the PCT</p>
V-2	National Patent (other kinds of protection or treatment, if any, are specified between parentheses after the designation(s) concerned)	<p>AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH&amp;LI CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW</p>
V-5	Precautionary Designation Statement In addition to the designations made under items V-1, V-2 and V-3, the applicant also makes under Rule 4.9(b) all designations which would be permitted under the PCT except any designation(s) of the State(s) indicated under item V-6 below. The applicant declares that those additional designations are subject to confirmation; and that any designation which is not confirmed before the expiration of 15 months from the priority date is to be regarded as withdrawn by the applicant at the expiration of that time limit.	
V-6	Exclusion(s) from precautionary designations	NONE
VI-1	Priority claim of earlier national application Filing date	04 September 1999 (04.09.1999)
VI-1-1	Number	9920872.0
VI-1-2	Country	GB
VII-1	International Searching Authority Chosen	European Patent Office (EPO) (ISA/EP)

## PCT REQUEST

Original (for SUBMISSION) - printed on 15.08.2000 04:08:03 PM

PU3517

VIII	Check list	number of sheets	electronic file(s) attached
VIII-1	Request	5	-
VIII-2	Description	403	-
VIII-3	Claims	26	-
VIII-4	Abstract	1	pu3517_abstract.txt
VIII-5	Drawings	0	-
VIII-7	<b>TOTAL</b>	<b>435</b>	
VIII-8	Accompanying items	paper document(s) attached	
VIII-9	Fee calculation sheet	✓	
VIII-10	Separate signed power of attorney	-	
VIII-16	PCT-EASY diskette	-	
VIII-18	Figure of the drawings which should accompany the abstract		
VIII-19	Language of filing of the international application	English	
IX-1	Signature of applicant or agent		
IX-1-1	Name (LAST, First)	CRAWLEY, Karen	

## FOR RECEIVING OFFICE USE ONLY

10-1	Date of actual receipt of the purported international application	31 AUG 2000	( 31 08. 2000 )
10-2	Drawings:		
10-2-1	Received		
10-2-2	Not received		
10-3	Corrected date of actual receipt due to later but timely received papers or drawings completing the purported international application		
10-4	Date of timely receipt of the required corrections under PCT Article 11(2)		
10-5	International Searching Authority	ISA/EP	
10-6	Transmittal of search copy delayed until search fee is paid		

## FOR INTERNATIONAL BUREAU USE ONLY

11-1	Date of receipt of the record copy by the International Bureau	
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## PCT/INT'L COOPERATION TREATY

PCT

REC'D 27 NOV 2001

## INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

Applicant's or agent's file reference PU3517	FOR FURTHER ACTION	See Notification of Transmittal of International Preliminary Examination Report (Form PCT/IPEA/416)
International application No. PCT/EP00/08487	International filing date (day/month/year) 31/08/2000	Priority date (day/month/year) 04/09/1999
International Patent Classification (IPC) or national classification and IPC C07D277/24		
Applicant GLAXO GROUP LIMITED et al.		
<p>1. This international preliminary examination report has been prepared by this International Preliminary Examining Authority and is transmitted to the applicant according to Article 36.</p> <p>2. This REPORT consists of a total of 7 sheets, including this cover sheet.</p> <p><input type="checkbox"/> This report is also accompanied by ANNEXES, i.e. sheets of the description, claims and/or drawings which have been amended and are the basis for this report and/or sheets containing rectifications made before this Authority (see Rule 70.16 and Section 607 of the Administrative Instructions under the PCT).</p> <p>These annexes consist of a total of sheets.</p> <p>3. This report contains indications relating to the following items:</p> <ul style="list-style-type: none"> <li>I <input checked="" type="checkbox"/> Basis of the report</li> <li>II <input type="checkbox"/> Priority</li> <li>III <input checked="" type="checkbox"/> Non-establishment of opinion with regard to novelty, inventive step and industrial applicability</li> <li>IV <input type="checkbox"/> Lack of unity of invention</li> <li>V <input checked="" type="checkbox"/> Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement</li> <li>VI <input type="checkbox"/> Certain documents cited</li> <li>VII <input type="checkbox"/> Certain defects in the international application</li> <li>VIII <input type="checkbox"/> Certain observations on the international application</li> </ul>		

Date of submission of the demand 23/03/2001	Date of completion of this report 22.11.2001
Name and mailing address of the international preliminary examining authority: European Patent Office D-80298 Munich Tel. +49 89 2399 - 0 Tx: 523656 epmu d Fax: +49 89 2399 - 4465	Authorized officer Usuelli, A Telephone No. +49 89 2399 7366



INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT

International application No. PCT/EP00/08487

I. Basis of the report

1. With regard to the elements of the international application (*Replacement sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to this report since they do not contain amendments (Rules 70.16 and 70.17)*):

Description, pages:

1-403 as originally filed

Claims, No.:

1-37 as originally filed

2. With regard to the language, all the elements marked above were available or furnished to this Authority in the language in which the international application was filed, unless otherwise indicated under this item.

These elements were available or furnished to this Authority in the following language: , which is:

- the language of a translation furnished for the purposes of the international search (under Rule 23.1(b)).
- the language of publication of the international application (under Rule 48.3(b)).
- the language of a translation furnished for the purposes of international preliminary examination (under Rule 55.2 and/or 55.3).

3. With regard to any nucleotide and/or amino acid sequence disclosed in the international application, the international preliminary examination was carried out on the basis of the sequence listing:

- contained in the international application in written form.
- filed together with the international application in computer readable form.
- furnished subsequently to this Authority in written form.
- furnished subsequently to this Authority in computer readable form.
- The statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.
- The statement that the information recorded in computer readable form is identical to the written sequence listing has been furnished.

4. The amendments have resulted in the cancellation of:

- the description, pages:
- the claims, Nos.:
- the drawings, sheets:

5.  This report has been established as if (some of) the amendments had not been made, since they have been considered to go beyond the disclosure as filed (Rule 70.2(c)):

# INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/EP00/08487

*(Any replacement sheet containing such amendments must be referred to under item 1 and annexed to this report.)*

## 6. Additional observations, if necessary:

### III. Non-establishment of opinion with regard to novelty, inventive step and industrial applicability

1. The questions whether the claimed invention appears to be novel, to involve an inventive step (to be non-obvious), or to be industrially applicable have not been examined in respect of:
  - the entire international application.
  - claims Nos. 27-30 (industrial applicability).

because:

- the said international application, or the said claims Nos. 27-30 relate to the following subject matter which does not require an international preliminary examination (*specify*):  
**see separate sheet**
- the description, claims or drawings (*indicate particular elements below*) or said claims Nos. are so unclear that no meaningful opinion could be formed (*specify*):
- the claims, or said claims Nos. are so inadequately supported by the description that no meaningful opinion could be formed.
- no international search report has been established for the said claims Nos.:-

2. A meaningful international preliminary examination cannot be carried out due to the failure of the nucleotide and/or amino acid sequence listing to comply with the standard provided for in Annex C of the Administrative Instructions:
  - the written form has not been furnished or does not comply with the standard.
  - the computer readable form has not been furnished or does not comply with the standard.

### V. Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

#### 1. Statement

Novelty (N)	Yes:	Claims 2-7, 9-37
	No:	Claims 1, 8
Inventive step (IS)	Yes:	Claims
	No:	Claims 1-37
Industrial applicability (IA)	Yes:	Claims 1-26, 31-37

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT**

International application No. PCT/EP00/08487

No: Claims

**2. Citations and explanations  
see separate sheet**

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT - SEPARATE SHEET**

International application No. PCT/EP00/08487

**Re Item III**

**Non-establishment of opinion**

Claims 27-30 relate to subject matter considered by this Authority to be covered by the provisions of Rule 67.1 (iv) PCT. Consequently, no opinion will be formulated with respect to the industrial applicability of the subject matter of this claim, cf. Article 34(4)(a)(i) PCT.

**Re Item V**

**Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement**

1- In the present Written Opinion reference is made to the following documents:

d1: J. MED. CHEM. vol. 38, no. 10, 1995, pages 1657-1665,

d2: FR 1 552 793 A

d3: US 4 883 815 A

d4: US 4 207 234 A

d5: J. MED. CHEM. (1985), 28(6), pages 699-707.

d6: JUSTUS LIEBIGS ANN. CHEM. (1968), 712, pages 73-8

d7: SCI. PHARM. (1971), 39(4), pages 267-92

**2- Novelty**

2.1- The compound of formula 15 disclosed in scheme II of d5, in which X=4-NO<sub>2</sub> and Y=2-Cl (cf. page 700 Method B, in conjunction with page 699, line 8 of paragraph "Chemistry"), is encompassed by present claims 1 and 8. The compound 15 in which X=Cl and Y=2-F, is excluded from these claims for the effect of the disclaimers. Accordingly, claims 1 and 8 do not comply with requirements of Art. 33(2) PCT.

2.2- Some compounds disclosed in d1, d2, d4, d6 and d7 contain a benzoyl group and an amide moiety attached by a linking group. These compounds differ from the compounds of the present application in that the groups attached to the nitrogen of the amide radical differ from the present groups R<sub>3</sub>/R<sub>4</sub>.

Compounds of d3 differ from the present compounds in that they lack the moiety - COR1.

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT - SEPARATE SHEET**

International application No. PCT/EP00/08487

**3- Inventive step**

3.1- D1 discloses a class of benzophenone derivatives acting as inhibitors of HIV-1 reverse transcriptase. This document is regarded as the closest state of the art. The technical problem can be seen in the provision of further inhibitors of HIV reverse transcriptase.

Said problem has been allegedly solved by present compounds of formulae (I) and (IV) (compounds of formulae IA, IB, IC, ID, II and III appear encompassed by formula I). For the purpose of assessing the inventive step during the International Preliminary Examination it is assumed that these compounds indeed act as HIV inhibitors, i.e. solve the above defined technical problem.

3.2- The compounds of Tables 1 and 2 of d1 differ from the compounds of the invention only for the nature of the group R attached to the phenyl ring of the aniline moiety. In the compounds of formula 1a-1I said R group is a radical of formula  $-O(CH_2)_2NEt_2$  or a hydrogen. In the corresponding position the compounds of the invention can be substituted, for instance, by a  $C_1-C_8$ alkylamino radical or by small group such as hydroxy and halogen.

Due to this structural similarity with the HIV-inhibitors disclosed in d1, it appears that the skilled person would have expected similar properties also from the present compounds.

Thus, the presence of an inventive step should be demonstrated preferably by means of comparative tests, e.g. by showing a quantitative superiority or unexpected effects vis-à-vis the closest prior art.

It is observed that the Applicant indicates in the present description (page 1 lines 30-33) that the compounds of d1 induce "resistance virus and were inactive against a common resistance strain". Considering the close similarity between the compounds of the invention and the compound of d1 and the broad scope of the formulae (I) and (IV), it should be credible from the comparative tests that any advantage over the compounds of d1 is provided by substantially all the claimed compounds.

The above considerations apply also to those compounds containing an heterocyclic moiety instead of a phenyl ring (i.e. compounds of claims 10-13 and 21). In fact the broad definition of heterocyclic group (cf. page 28 of the description) encompasses also rings known as bioisosters of the phenyl group.

Accordingly it appears that there are no inventive contributions in the introduction of this feature.

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT - SEPARATE SHEET**

International application No. PCT/EP00/08487

**4- Industrial applicability**

For the assessment of the present claims 27-30 on the question whether they are industrially applicable, no unified criteria exist in the PCT Contracting States. The patentability can also be dependent upon the formulation of the claims. The EPO, for example, does not recognize as industrially applicable the subject-matter of claims to the use of a compound in medical treatment, but may allow, however, claims to a known compound for first use in medical treatment and the use of such a compound for the manufacture of a medicament for a new medical treatment.

## PATENT COOPERATION TREATY

From the  
INTERNATIONAL PRELIMINARY EXAMINING AUTHORITY:

To:

CRAWLEY, Karen  
GLAXOSMITHKLINE  
Corporate Intellectual Property  
Two New Horizons Court  
Brentford  
Middlesex TW8 9EP  
GRANDE BRETAGNE

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25 NOV 2001

NEW HORIZONS COURT

NOTIFICATION OF TRANSMITTAL OF  
THE INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT

(PCT Rule 71.1)

Action Date

PCT

06 DEC 2001

RE

RE

Date of mailing  
(day/month/year)

22.11.2001

Applicant's or agent's file reference  
PU3517

## IMPORTANT NOTIFICATION

International application No.  
PCT/EP00/08487International filing date (day/month/year)  
31/08/2000Priority date (day/month/year)  
04/09/1999

Applicant

GLAXO GROUP LIMITED et al.

1. The applicant is hereby notified that this International Preliminary Examining Authority transmits herewith the international preliminary examination report and its annexes, if any, established on the international application.
2. A copy of the report and its annexes, if any, is being transmitted to the International Bureau for communication to all the elected Offices.
3. Where required by any of the elected Offices, the International Bureau will prepare an English translation of the report (but not of any annexes) and will transmit such translation to those Offices.

## 4. REMINDER

The applicant must enter the national phase before each elected Office by performing certain acts (filing translations and paying national fees) within 30 months from the priority date (or later in some Offices) (Article 39(1)) (see also the reminder sent by the International Bureau with Form PCT/IB/301).

Where a translation of the international application must be furnished to an elected Office, that translation must contain a translation of any annexes to the international preliminary examination report. It is the applicant's responsibility to prepare and furnish such translation directly to each elected Office concerned.

For further details on the applicable time limits and requirements of the elected Offices, see Volume II of the PCT Applicant's Guide.

Name and mailing address of the IPEA/

European Patent Office  
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Tel. +49 89 2399 - 0 Tx: 523656 epmu d  
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Authorized officer

Ambrona, J.R.

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# PATENT COOPERATION TREATY

## PCT

### INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

Applicant's or agent's file reference PU3517	FOR FURTHER ACTION		See Notification of Transmittal of International Preliminary Examination Report (Form PCT/IPEA/416)
International application No. PCT/EP00/08487	International filing date (day/month/year) 31/08/2000	Priority date (day/month/year) 04/09/1999	
International Patent Classification (IPC) or national classification and IPC C07D277/24			
Applicant GLAXO GROUP LIMITED et al.			

<p>1. This international preliminary examination report has been prepared by this International Preliminary Examining Authority and is transmitted to the applicant according to Article 36.</p> <p>2. This REPORT consists of a total of 7 sheets, including this cover sheet.</p> <p><input type="checkbox"/> This report is also accompanied by ANNEXES, i.e. sheets of the description, claims and/or drawings which have been amended and are the basis for this report and/or sheets containing rectifications made before this Authority (see Rule 70.16 and Section 607 of the Administrative Instructions under the PCT).</p> <p>These annexes consist of a total of sheets.</p>	
<p>3. This report contains indications relating to the following items:</p> <ul style="list-style-type: none"> <li>I <input checked="" type="checkbox"/> Basis of the report</li> <li>II <input type="checkbox"/> Priority</li> <li>III <input checked="" type="checkbox"/> Non-establishment of opinion with regard to novelty, inventive step and industrial applicability</li> <li>IV <input type="checkbox"/> Lack of unity of invention</li> <li>V <input checked="" type="checkbox"/> Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement</li> <li>VI <input type="checkbox"/> Certain documents cited</li> <li>VII <input type="checkbox"/> Certain defects in the international application</li> <li>VIII <input type="checkbox"/> Certain observations on the international application</li> </ul>	

Date of submission of the demand 23/03/2001	Date of completion of this report 22.11.2001
Name and mailing address of the international preliminary examining authority: European Patent Office D-80298 Munich Tel. +49 89 2399 - 0 Tx: 523656 epmu d Fax: +49 89 2399 - 4465	Authorized officer Usuelli, A Telephone No. +49 89 2399 7366



**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT**

International application No. PCT/EP00/08487

**I. Basis of the report**

1. With regard to the elements of the international application (Replacement sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to this report since they do not contain amendments (Rules 70.16 and 70.17)):

**Description, pages:**

1-403 as originally filed

**Claims, No.:**

1-37 as originally filed

2. With regard to the language, all the elements marked above were available or furnished to this Authority in the language in which the international application was filed, unless otherwise indicated under this item.

These elements were available or furnished to this Authority in the following language: , which is:

- the language of a translation furnished for the purposes of the international search (under Rule 23.1(b)).
- the language of publication of the international application (under Rule 48.3(b)).
- the language of a translation furnished for the purposes of international preliminary examination (under Rule 55.2 and/or 55.3).

3. With regard to any nucleotide and/or amino acid sequence disclosed in the international application, the international preliminary examination was carried out on the basis of the sequence listing:

- contained in the international application in written form.
- filed together with the international application in computer readable form.
- furnished subsequently to this Authority in written form.
- furnished subsequently to this Authority in computer readable form.
- The statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.
- The statement that the information recorded in computer readable form is identical to the written sequence listing has been furnished.

4. The amendments have resulted in the cancellation of:

- the description, pages:
- the claims, Nos.:
- the drawings, sheets:

5.  This report has been established as if (some of) the amendments had not been made, since they have been considered to go beyond the disclosure as filed (Rule 70.2(c)):

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT**

International application No. PCT/EP00/08487

*(Any replacement sheet containing such amendments must be referred to under item 1 and annexed to this report.)*

**6. Additional observations, if necessary:**

**III. Non-establishment of opinion with regard to novelty, inventive step and industrial applicability**

1. The questions whether the claimed invention appears to be novel, to involve an inventive step (to be non-obvious), or to be industrially applicable have not been examined in respect of:

- the entire international application.
- claims Nos. 27-30 (industrial applicability).

because:

- the said international application, or the said claims Nos. 27-30 relate to the following subject matter which does not require an international preliminary examination (specify):  
see separate sheet
- the description, claims or drawings (*indicate particular elements below*) or said claims Nos. are so unclear that no meaningful opinion could be formed (specify):
- the claims, or said claims Nos. are so inadequately supported by the description that no meaningful opinion could be formed.
- no international search report has been established for the said claims Nos. .

2. A meaningful international preliminary examination cannot be carried out due to the failure of the nucleotide and/or amino acid sequence listing to comply with the standard provided for in Annex C of the Administrative Instructions:

- the written form has not been furnished or does not comply with the standard.
- the computer readable form has not been furnished or does not comply with the standard.

**V. Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement**

**1. Statement**

Novelty (N)	Yes:	Claims 2-7, 9-37
	No:	Claims 1, 8
Inventive step (IS)	Yes:	Claims
	No:	Claims 1-37
Industrial applicability (IA)	Yes:	Claims 1-26, 31-37

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT**

International application No. PCT/EP00/08487

No: Claims

2. Citations and explanations  
see separate sheet

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT - SEPARATE SHEET**

International application No.

PCT/EP00/08487

**Re Item III**

**Non-establishment of opinion**

Claims 27-30 relate to subject matter considered by this Authority to be covered by the provisions of Rule 67.1 (iv) PCT. Consequently, no opinion will be formulated with respect to the industrial applicability of the subject matter of this claim, cf. Article 34(4)(a)(i) PCT.

**Re Item V**

**Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement**

1- In the present Written Opinion reference is made to the following documents:

d1: J. MED. CHEM. vol. 38, no. 10, 1995, pages 1657-1665,

d2: FR 1 552 793 A

d3: US 4 883 815 A

d4: US 4 207 234 A

d5: J. MED. CHEM. (1985), 28(6), pages 699-707.

d6: JUSTUS LIEBIGS ANN. CHEM. (1968), 712, pages 73-8

d7: SCI. PHARM. (1971), 39(4), pages 267-92

2- Novelty

2.1- The compound of formula 15 disclosed in scheme II of d5, in which X=4-NO<sub>2</sub> and Y=2-Cl (cf. page 700 Method B, in conjunction with page 699, line 8 of paragraph "Chemistry"), is encompassed by present claims 1 and 8. The compound 15 in which X=Cl and Y=2-F, is excluded from these claims for the effect of the disclaimers. Accordingly, claims 1 and 8 do not comply with requirements of Art. 33(2) PCT.

2.2- Some compounds disclosed in d1, d2, d4, d6 and d7 contain a benzoyl group and an amide moiety attached by a linking group. These compounds differ from the compounds of the present application in that the groups attached to the nitrogen of the amide radical differ from the present groups R<sub>3</sub>/R<sub>4</sub>.

Compounds of d3 differ from the present compounds in that they lack the moiety - COR1.

INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT - SEPARATE SHEET

International application No.

PCT/EP00/08487

3- Inventive step

3.1- D1 discloses a class of benzophenone derivatives acting as inhibitors of HIV-1 reverse transcriptase. This document is regarded as the closest state of the art. The technical problem can be seen in the provision of further inhibitors of HIV reverse transcriptase.

Said problem has been allegedly solved by present compounds of formulae (I) and (IV) (compounds of formulae IA, IB, IC, ID, II and III appear encompassed by formula I). For the purpose of assessing the inventive step during the International Preliminary Examination it is assumed that these compounds indeed act as HIV inhibitors, i.e. solve the above defined technical problem.

3.2- The compounds of Tables 1 and 2 of d1 differ from the compounds of the invention only for the nature of the group R attached to the phenyl ring of the aniline moiety. In the compounds of formula 1a-1I said R group is a radical of formula  $-\text{O}(\text{CH}_2)_2\text{NEt}_2$  or a hydrogen. In the corresponding position the compounds of the invention can be substituted, for instance, by a  $\text{C}_1\text{-C}_6$ alkylamino radical or by small groups such as hydroxy and halogen.

Due to this structural similarity with the HIV-inhibitors disclosed in d1, it appears that the skilled person would have expected similar properties also from the present compounds.

Thus, the presence of an inventive step should be demonstrated preferably by means of comparative tests, e.g. by showing a quantitative superiority or unexpected effects vis-à-vis the closest prior art.

It is observed that the Applicant indicates in the present description (page 1 lines 30-33) that the compounds of d1 induce "resistance virus and were inactive against a common resistance strain". Considering the close similarity between the compounds of the invention and the compound of d1 and the broad scope of the formulae (I) and (IV), it should be credible from the comparative tests that any advantage over the compounds of d1 is provided by substantially all the claimed compounds.

The above considerations apply also to those compounds containing a heterocyclic moiety instead of a phenyl ring (i.e. compounds of claims 10-13 and 21). In fact the broad definition of heterocyclic group (cf. page 28 of the description) encompasses also rings known as bioisosters of the phenyl group.

Accordingly it appears that there are no inventive contributions in the introduction of this feature.

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT - SEPARATE SHEET**

International application No.

PCT/EP00/08487

**4- Industrial applicability**

For the assessment of the present claims 27-30 on the question whether they are industrially applicable, no unified criteria exist in the PCT Contracting States. The patentability can also be dependent upon the formulation of the claims. The EPO, for example, does not recognize as industrially applicable the subject-matter of claims to the use of a compound in medical treatment, but may allow, however, claims to a known compound for first use in medical treatment and the use of such a compound for the manufacture of a medicament for a new medical treatment.

TO:  
PCT Examination  
International Bureau of WIPO  
34 Chimin des Colombettes  
1211 Geneva 20  
Switzerland

Fax: 011 41 22 740 1435

Direction to PCT  
Request before  
expiration of 30th  
Month

Applicant's File Reference PU3517WO	Applicant Glaxo Group Limited
International Application No. PCT/EP00/08487	International Filing Date: 31 August 2000
30th Month Deadline: 4 March 2002	Title: Benzophenones as Inhibitors of Reverse Transcriptase

Please make the following corrections to PCT Request :

Please change the name of inventor Clarence Webster ANDREWS to Clarence Webster ANDREWS III

Box III-1 Please change the address of inventor Clarence Webster ANDREWS, III from , Glaxo Wellcome Inc., Five Moore Drive, PO Box 13398, Research Triangle Park, NC 27709, US

Change to

GlaxoSmithKline, Five Moore Drive, PO Box 13398, Research Triangle Park, NC 27709, US

Box III-2 Please change the address of inventor Joseph Howing CHAN from Glaxo Wellcome Inc., Five Moore Drive, PO Box 13398, Research Triangle Park, NC 27709, US

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Box III-3 Please change the address of inventor George Andrew FREEMAN from Glaxo Wellcome Inc., Five Moore Drive, PO Box 13398, Research Triangle Park, NC 27709, US

Change to

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Box III-5 Please change the address of inventor Karen Rene ROMINES from Glaxo Wellcome Inc., Five Moore Drive, PO Box 13398, Research Triangle Park, NC 27709, US

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GlaxoSmithKline, Five Moore Drive, PO Box 13398, Research Triangle Park, NC 27709, US

Box III-6 Please change the address of inventor Jeffrey H. TIDWELL from Glaxo Wellcome Inc., Five Moore Drive, PO Box 13398, Research Triangle Park, NC 27709, US

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Box III-7 Please change the address of inventor Pascal Maurice Charles PLANETTI from Laboratoire Glaxo Wellcome, Centre de Recherches, Z.A. de Couraboeuf, 25, avenue de Québec, F-91940 Les Ulis, France

Change to

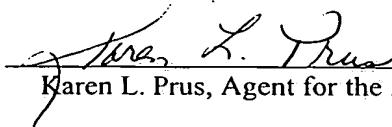
Laboratoire GlaxoSmithKline, Centre de Recherches, Z.A. de Couraboeuf, 25, avenue de Québec, F-91940 Les Ulis, France

Please confirm receipt of this Request to Change PCT Request for 30th Month to fax 919-483-7988.

Please send PCT/IB/306 to: David J. Levy, VP, GlaxoSmithKline, Corporate Intellectual Property Department, Five Moore Drive, PO Box 13398, Research Triangle Park, NC 27709

If there should be questions, please call (919) 483-2252.

Thank you.  
Sincerely,

 2/18/02  
Karen L. Prus, Agent for the Applicant

**FAXE**  
2-18-02